U.S. Serial No.: 09/587,111 Filed: June 2, 2000 Title: NOVEL MEMBERS OF THE CAPSAICIN/VANILLOID RECEPTOR FAMILY OF PROTEINS AND USES THEREOF

Agent: Mario Cloutier

Docket No.: MPI98-093P2RCP3DV1AM

1/37

NumanVRl gene with translation of open reading frame

Input file Fchrb87a6.seq; Output File Fchrb87a6.tra Sequence length 3909

GTGAGCGCAACGCACTGCGGGCAGTGAGCGCAACGCACTGCGGGCAGTGAGCGCAACGCACTGCGGGCAGTGAGCGCAA CGCACTGCGGGCAGTGAGCGCAACGCACTGCGGGCAGTGAGCGCAACGCACTGCGGGCAGTGAGCGCAACGCACTGCGG GCAGTGAGCGCAACGCACTTGCGGGCAGTGAGCGCAACGCACTGCGGGCAGTGAGCGCAACGCACTGCGGGCAGTGAGC GCAACGCACTGCGGGCAGTGAGCGCAACGCACTGCGGGCAGTGAGCGCAACGCACTGCGGGCAGTGAGCGCAACGCACT TTTATGCTTCCGGCTCGTATGTTGTGTGGAATTGTGAGCGGATAACAATTTCACACAGGAAACAGCTATGACCATGATT ACGCCAAGCTCTAATACGACTCACTATAGGGAAAGCTGGTACGCCTGCAGGTACCGGTCCGGAATTCCCGGGTCGACCC ACGCGTCCGAAAACACACCTCTCTGCTGTGGGAAGACTGTGCAATGGCACAGCCGCAGAGCTTGGTTTGGGAGGTTGAA GTGCTCTGGGGAGAATTCGTAGATCATCCTCAGAAAAGCCTTGCCCTGGTGTTCTACCAGAAAAACGTCTCCCAATCAC CCAGAAAAGCTGTCCACAGTAGTCCCCCCTTATCCACGGGTGTCACTTTCCATGGGTTCAGTTATTTGCGGTCAACCAC GGTCTGCCAATATTAAATGGAAAATTCTTCAAACAGTTCCCCAAGTTTTCCCTTGTGCATTGTTCTGAGCAGTGTGATGA AGAGTCTCTGCCGTGCCATCTGGGATGCAAACCGTCCCTGTGTCCCCACGTCCAGGCCGTAGATGCTCCCCGCCGGTC **AGTCACTTAGTCGTCAGATCGCCCGTCCTGGTATCACAGTGCTTCTGTTCAGGTTGCACACTGGGCCACAGAGGATCCA** S D L G T 18 GCAAGG ATG AAG AAA TGG AGC AGC ACA GAC TTG GGG ACA GCT GCG GAC CCA CTC CAA AAG 54 D P G D P P P P 38 L D N S R A GAC ACC TGC CCA GAC CCC CTG GAT GGA GAC CCT AAC TCC AGG CCA CCT CCA GCC AAG CCC 114 A K R T RL F G K G D CAG CTC CCC ACG GCC AAG AGC CGC ACC CGG CTC TTT GGG AAG GGT GAC TCG GAG GAG GCT 174 E S H E G E D L TTC CCG GTG GAT TGC CCC CAC GAG GAA GGT GAG TTG GAC TCC TGC CCG ACC ATC ACA GTC 234 G 98 R P G D P T G A R L L Τ T

AGC CCT GTT ATC ACC ATC CAG AGG CCA GGA GAC GGC CCC ACC GGT GCC AGG CTG CTG TCC 294

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S T E K T L RLYD A CAG GAC TCT GTC GCC GCC AGC ACC GAG AAG ACC CTC AGG CTC TAT GAT CGC AGG AGT ATC 354 ONNCODLESL L L F L TTT GAA GCC GTT GCT CAG AAT AAC TGC CAG GAT CTG GAG AGC CTG CTG CTC CTG CAG 414 K S K K H L T D N E F K D P E T G K T C AAG AGC AAG AAG CAC CTC ACA GAC AAC GAG TTC AAA GAC CCT GAG ACA GGG AAG ACC TGT G 178 AMLNLHD Q N T CTG CTG AAA GCC ATG CTC AAC CTG CAC GAC GGA CAG AAC ACC ACC ATC CCC CTG CTC CTG 198 EIARQTDSLKELVNA T D S S Y GAG ATC GCG CGG CAA ACG GAC AGC CTG AAG GAG CTT GTC AAC GCC AGC TAC ACG GAC AGC 594 218 YYKGQTALHIAIERR N M TAC TAC AAG GGC CAG ACA GCA CTG CAC ATC GCC ATC GAG AGA CGC AAC ATG GCC CTG GTG V E N G A D V Q 238 A A A H ACC CTC CTG GTG GAG AAC GGA GCA GAC GTC CAG GCT GCG GCC CAT GGG GAC TTC TTT AAG 714 K T K G R P G F Y F G E L P L S L A A C AAA ACC AAA GGG CGG CCT GGA TTC TAC TTC GGT GAA CTG CCC CTG TCC CTG GCC GCG TGC 774 278 IVKF LLON ACC AAC CAG CTG GGC ATC GTG AAG TTC CTG CTG CAG AAC TCC TGG CAG ACG GCC GAC ATC 834 298 S A R D V G T .V L HALVEVADN N AGC GCC AGG GAC TCG GTG GGC AAC ACG GTG CTG CAC GCC CTG GTG GAG GTG GCC GAC AAC 894 TADNTKFVT SMYNEILML ACG GCC GAC AAC ACG AAG TTT GTG ACG AGC ATG TAC AAT GAG ATT CTG ATG CTG GGG GCC L K L E E L TNKK G M AAA CTG CAC CCG ACG CTG AAG CTG GAG GAG CTC ACC AAC AAG AAG GGA ATG ACG CCG CTG 1014 A A G T G K I G V L A Y I L Q R E I GCT CTG GCA GCT GGG ACC GGG AAG ATC GGG GTC TTG GCC TAT ATT CTC CAG CGG GAG ATC 1074 CRHLSRKFTEW A CAG GAG CCC GAG TGC AGG CAC CTG TCC AGG AAG TTC ACC GAG TGG GCC TAC GGG CCC GTG 1134 C E K N L Y D L SCID T CAC TCC TCG CTG TAC GAC CTG TCC TGC ATC GAC ACC TGC GAG AAG AAC TCG GTG CTG GAG 1194 Y S S S E T P N R H D M L L GTG ATC GCC TAC AGC AGC AGC GAG ACC CCT AAT CGC CAC GAC ATG CTC TTG GTG GAG CCG 1254

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L Q D K W D R F V K R I F Y F N CTG AAC CGA CTC CTG CAG GAC AAG TGG GAC AGA TTC GTC AAG CGC ATC TTC TAC TTC AAC 1314 LYMI I F T M A A Y Y TTC CTG GTC TAC TGC CTG TAC ATG ATC ATC TTC ACC ATG GCT GCC TAC TAC AGG CCC GTG 1374 P P F K M E KIG D Y F R V 478 GAT GGC TTG CCT CCC TTT AAG ATG GAA ARA ATT GGA GAC TAT TTC CGA GTT ACT GGA GAG 1434 GGVYFFFRG ATC CTG TCT GTG TTA GGA GGA GTC TAC TTC TTT TTC CGA GGG ATT CAG TAT TTC CTG CAG 1494 K T L F Y V D S S E AGG CGG CCG TCG ATG AAG ACC CTG TTT GTG GAC AGC TAC AGT GAG ATG CTT TTC TTT CTG 1554 LFMLATVVLYF S HLK E Y V 538 CAG TCA CTG TTC ATG CTG GCC ACC GTG GTG CTG TAC TTC AGC CAC CTC AAG GAG TAT GTG 1614 F SLAL G W T N M L Y Y GCT TCC ATG GTA TTC TCC CTG GCC TTG GGC TGG ACC AAC ATG CTC TAC TAC ACC CGC GGT 1674 GIYAVM Ι E K M Ι L R D 578 L TTC CAG CAG ATG GGC ATC TAT GCC GTC ATG ATA GAG AAG ATG ATC CTG AGA GAC CTG TGC 1734 VYIVFLFGF S A CGT TTC ATG TTT GTC TAC ATC GTC TTC TTG TTC GGG TTT TCC ACA GCG GTG GTG ACG CTG 1794 D S L P S E ATT GAA GAC GGG AAG AAT GAC TCC CTG CCG TCT GAG TCC ACG TCG CAC AGG TGG CGG GGG 1854 P A C R P P D S S Y N S L Y S T CLE 638 CCT GCC TGC AGG CCC CCC GAT AGC TCC TAC AAC AGC CTG TAC TCC ACC TGC CTG GAG CTG 1914 T I G M KF G D L E F T E Y N D F K TTC AAG TTC ACC ATC GGC ATG GGC GAC CTG GAG TTC ACT GAG AAC TAT GAC TTC AAG GCT 1974 I L L L A Y V L Ι I T Y I L L L N GTC TTC ATC ATC CTG CTG CTG GCC TAT GTA ATT CTC ACC TAC ATC CTC CTG CTC AAC ATG 2034 G E T V N K I A Q K N L M S 698 E CTC ATC GCC CTC ATG GGT GAG ACT GTC AAC AAG ATC GCA CAG GAG AGC AAG AAC ATC TGG 2094 Ι d. I L D T E K S F L K С M A AAG CTG CAG AGA GCC ATC ACC ATC CTG GAC ACG GAG AAG AGC TTC CTT AAG TGC ATG AGG 2154

Applicant: Rory A. J. Curtis
U.S. Serial No.: 09/587,111 Filed: June 2, 2000
Title: NOVEL MEMBERS OF THE CAPSAICIN/VANILLOID
RECEPTOR FAMILY OF PROTEINS AND USES THEREOF
Agent: Mario Cloutier

Docket No.: MPI98-093P2RCP3DV1AM

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																				738 2214
																				758 2274
																				778 2334
																				798 2394
																				818 2454
																				838 2514
	* TGA																			840 2520
GGA	CGTC	ACGC	AGAC	AGCA	CTGT	CAACI	CTG	GCC:	rtag(GAGA	CCCC	3TTG(CAC	GGGG	GCT(GCTG/	AGGGI	AACA(CAG	

Fig. 1D

market and the second

Filed: June 2, 2000 Title: NOVEL MEMBERS OF THE CAPSAICIN/VANILLOID RECEPTOR FAMILY OF PROTEINS AND USES THEREOF

Agent: Mario Cloutier

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Full-length human VR2

Input file Flh21e11.seq; Output File Flh21e11.tra Sequence length 2809

GGCTAGCCTGTCCTGACAGGGGAGAGTTAAGCTCCCGTTCTCCACCGTGCCGGCTGGCCAGGTGGGCTGAGGGTGACCG GGCAGCCCCTCCCGGCTTCACTTCCTCCCGCAGCCCCTGCTACTGAGAAGCTCCGGGATCCCAGCAGCCGCCACGCCCT <u>ርድኖሮሞና እርድሮሞርምርድርድምናር እርሞር እርድርር እርድርር እርድርር እርድሞርድር እርድ እርድር እርድ እርድር እርድ እርድርር እርድ እርድርርር እርድርርር እርድርርር</u> እርድር

GGCCTCAGCCTGCGGGGCTCCAGTCAGGCCAACACCGACGCGCAGCTGGGAAGACAGGACCCCTTGACATCTCCATC																	
TGC	CAG	\GGT(CTG	CTG	BACCO	BAGCI	\GCC1	rcct(CTC	TAG		T ACC					8 24
					T ACA												28 84
																R CGG	48 144
					I ATA											Q CAG	68 204
					F TTT											P CCC	88 264
																S TCG	108 324
					S TCC		_									K AAG	128 384
					C TGC											P CCT	148 444
					A GCC												168 504
					R AGG												188 564
-					C TGC												208 624
G GGT	E GAG		P													L CTC	228 684

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Agent: Mario Cloutier

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P A S L Q A T D S Q G N H CTG GAG AAC CCA CAC CAG CCC GCC AGC CTG CAG GCC ACT GAC TCC CAG GGC AAC ACA GTC 744 M I S D N S A E N I A L 268 CTG CAT GCC CTA GTG ATG ATC TCG GAC AAC TCA GCT GAG AAC ATT GCA CTG GTG ACC ACC 804 288 G L L Q A G A R L C P T V Q L E ATG TAT GAT GGG CTC CTC CAA GCT GGG GCC CGC CTC TGC CCT ACC GTG CAG CTT GAG GAC 864 LQDLTPLKLAAKE 308 ATC GCC AAC CTG CAG GAT CTC ACG CCT CTG AAG CTG GCC GCC AAG GAG GGC AAG ATC GAG 924 H ILOREF S G L SH L S R 328 ATT TTC AGG CAC ATC CTG CAG CGG GAG TTT TCA GGA CTG AGC CAC CTT TCC CGA AAG TTC 984 RVSLYDLA ACC GAG TGG TGC TAT GGG CCT GTC CGG GTG TCG CTG TAT GAC CTG GCT TCT GTG GAC AGC 1044 VLE IIAFHCKS TGT GAG GAG AAC TCA GTG CTG GAG ATC ATT GCC TTT CAT TGC AAG AGC CCG CAC CGA CAC 1104 V V L E P L N K L L Q A K W D L L I CGA ATG GTC GTT TTG GAG CCC CTG AAC AAA CTG CTG CAG GCG AAA TGG GAT CTG CTC ATC 1164 F F L N F L C N L I Y M F Ι F T CCC AAG TTC TTC AAC TTC CTG TGT AAT CTG ATC TAC ATG TTC ATC TTC ACC GCT GTT 1224 HQPTLKKQA A P H L K A E GCC TAC CAT CAG CCT ACC CTG AAG AAG CAG GCC GCC CCT CAC CTG AAA GCG GAG GTT GGA 1284 MLLT G H I L I L L G G IYLL AAC TCC ATG CTG ACG GGC CAC ATC CTT ATC CTG CTA GGG GGG ATC TAC CTC CTC GTG 1344 YFWR R H V Ι F GGC CAG CTG TGG TAC TTC TGG CGG CGC CAC GTG TTC ATC TGG ATC TCG TTC ATA GAC AGC 1404 L T V V EILFLFQAL S OVL TAC TTT GAA ATC CTC TTC CTG TTC CAG GCC CTG CTC ACA GTG GTG TCC CAG GTG CTG TGT 1464 EWYLPL L V A I V S A L L 508 TTC CTG GCC ATC GAG TGG TAC CTG CCC CTG CTT GTG TCT GCG CTG GTG CTG GGC TGG CTG 1524 G I Y LYYTRGF O H T V M Ι AAC CTG CTT TAC TAT ACA CGT GGC TTC CAG CAC ACA GGC ATC TAC AGT GTC ATG ATC CAG 1584 Applicant: Rory A. J. Curtis
U.S. Serial No.: 09/587,111 Filed: June 2, 2000
Title: NOVEL MEMBERS OF THE CAPSAICIN/VANILLOID
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Agent: Mario Cloutier

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K	V	I	L	R	D	L	L	R	F	L	L	I	Y	L	V	F	L	F	G	548
AAG	GTC	ATC	CTG	CGG	GAC	CTG	CTG	CGC	TTC	CTT	CTG	ATC	TAC	TTA	GTC	TTC	CTT	TTC	GGC	1644
																				568 1704
P	n	A	T	E	S	V	Q	P	M	E	G	Q	E	D	E	G	N	G	A	588
CCC	aat	GCC	ACA	GAG	TCA	GTG	CAG		ATG	GAG	GGA	CAG	GAG	GAC	GAG	GGC	AAC	GGG	GCC	1764
Q	Y	R	G	I	L	E	A	S	L	E	L	F	K	F	T	I	G	M	G	608
CAG	TAC	AGG	GGT	ATC	CTG	Gaa	GCC	TCC	TTG	GAG	CTC	TTC	AAA	TTC	ACC	ATC	GGC	ATG	GGC	1824
E	L	A	F	Q	E	Q	L	H	F	R	G	M	V	L	L	L	L	L	A	628
GAG	CTG	GCC	TTC	CAG	GAG	CAG	CTG	CAC	TTC	CGC	GGC	ATG	GTG	CTG	CTG	CTG	CTG	CTG	GCC	1884
																				648 1944
V	N	S	V	A	T	D	S	W	S	I	W	K	L	Q	K	A	I	S	V	668
GTC	AAC	AGT	GTC	GCC	ACT	GAC	AGC	TGG	AGC	ATC	TGG	AAG	CTG	CAG	AAA	GCC	ATC	TCT	GTC	2004
L	E	M	E	n	G	y	W	W	C	R	K	K	Q	R	A	G	V	M	L	688
CTG	GAG	ATG	GAG	aat	GGC	Tat	TGG	TGG	TGC	AGG	AAG	AAG	CAG	CGG	GCA	GGT	GTG	ATG	CTG	2064
T	V	G	T	K	P	D	G	S	P	D	E	R	W	C	F	R	v	E	E	708
ACC	GTT	GGC	ACT	AAG	CCA	GAT	GGC	AGC	CCG	GAT	GAG	CGC	TGG	TGC	TTC	AGG	G T G	GAG	GAG	2124
V	N	W	A	S	W	E	Q	T	L	P	T	L	C	E	D	P	S	G	A	728
GTG	AAC	TGG	GCT	TCA	TGG	GAG	CAG	ACG	CTG	CCT	ACG	CTG	TGT	GAG	GAC	CCG	TCA	GGG	GCA	2184
																				748 2244
																* TGA				765 2295

Fig. 2C

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Agent: Mario Cloutier

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Partial human VR2 alternate form

Input file frhob12c4.seg; Output File frhob12c4.tra Sequence length 1489

T C FYF 19 OKGOG G GC GGC CGC TTC TTC CAG AAG GGC CAA GGG ACT TGC TTT TAT TTC GGT GAG CTA CCC CTC 57 39 ACTK O W D V v S Y L L E N TCT TTG GCC GCT TGC ACC AAG CAG TGG GAT GTG GTA AGC TAC CTC CTG GAG AAC CCA CAC 117 A T D S Q G N CAG CCC GCC AGC CTG CAG GCC ACT GAC TCC CAG GGC AAC ACA GTC CTG CAT GCC CTA GTG 177 79 A E N Ι A L ATG ATC TCG GAC AAC TCA GCT GAG AAC ATT GCA CTG GTG ACC AGC ATG TAT GAT GGG CTC 237 99 L C P T V Q L E D CTC CAA GCT GGG GCC CGC CTC TGC CCT ACC GTG CAG CTT GAG GAC ATC CGC AAC CTG CAG 297 L L K E G K I E I F R 119 K A GAT CTC ACG CCT CTG AAG CTG GCC GCC AAG GAG GGC AAG ATC GAG ATT TTC AGG CAC ATC 357 R E F G L S H L S R K F T Ε 139 S CTG CAG CGG GAG TTT TCA GGA CTG AGC CAC CTT TCC CGA AAG TTC ACC GAG TGG TGC TAT 417 ٧ D С 159 L Y D T. A S GGG CCT GTC CGG GTG TCG CTG TAT GAC CTG GCT TCT GTG GAC AGC TGT GAG GAG AAC TCA 477 Ι A F H C K S P H R 179 Ι GTG CTG GAG ATC ATT GCC TTT CAT TGC AAG AGC CCG CAC CGA CAC CGA ATG GTC GTT TTG 537 L 0 A K W D L L Ι P F 199 GAG CCC CTG AAC AAA CTG CTG CAG GCG AAA TGG GAT CTG CTC ATC CCC AAG TTC TTC TTA 219 Y M F F V A Y I Ι A AAC TTC CTG TGT AAT CTG ATC TAC ATG TTC ATC TTC ACC GCT GTT GCC TAC CAT CAG CCT 657 A P H K E V G N 239 L A Q A ACC CTG AAG AAG CAG GCC GCC CCT CAC CTG AAA GCG GAG GTT GGA AAC TCC ATG CTG CTG 717 259 G G I Y L IL I L L 777 ACG GGC CAC ATC CTT ATC CTG CTA GGG GGG ATC TAC CTC CTC GTG GGC CAG CTG TGG TAC Y 279 F Ι W I S F Ι D S F E H TTC TGG CGG CGC CAC GTG TTC ATC TGG ATC TCG TTC ATA GAC AGC TAC TTT GAA ATC CTC 837 Applicant: Rory A. J. Curtis
U.S. Serial No.: 09/587,111 Filed: June 2, 2000
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QALLTVVSO LCFLAIE TTC CTG TTC CAG GCC CTG CTC ACA GTG GTG TCC CAG GTG CTG TGT TTC CTG GCC ATC GAG 897 PLLV S A L ٧ L 319 TGG TAC CTG CCC CTG CTT GTG TCT GCG CTG GTG CTG GGC TGG CTG AAC CTG CTT TAC TAT F Q H T G Ι Y S V M 339 Ι Q K ACA CGT GGC TTC CAG CAC ACA GGC ATC TAC AGT GTC ATG ATC CAG AAG AAA GCC ATC TCT 1017 m e N G Y C 359 W W R K K R GTC CTG GAG ATG GAG AAT GGC TAT TGG TGG TGC AGG AAG AAG CAG CGG GCA GGT GTG ATG 1077 GTKPDGS P D Ë R W C F R V E CTG ACC GTT GGC ACT AAG CCA GAT GGC AGC CCG GAT GAG CGC TGG TGC TTC AGG GTG GAG 1137 W E O T L P 399 GAG GTG AAC TGG GCT TCA TGG GAG CAG ACG CTG CCT ACG CTG TGT GAG GAC CCG TCA GGG 1197 R T L E N P L Α S K E D GCA GGT GTC CCT CGA ACT CTC GAG AAC CCT GTC CTG GCT TCC CCT CCC AAG GAG GAT GAG 1257 N Y V P V 437 GAT GGT GCC TCT GAG GAA AAC TAT GTG CCC GTC CAG CTC CTC CAG TCC AAC TGA 1311 GCGGACGCGTGGGTCGAC

Fig. 3B

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Agent: Mario Cloutier

Docket No.: MPI98-093P2RCP3DV1AM

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Partial rat VR2

Input file Flrxb147g11.seq; Output File Flrxb147g11.tra
Sequence length 1794

				S 1 CC GC									19 57
				N AAC							L CTG		39 117
				A GCT									59 177
				D GAC									79 237
L CTT	_		I ATC	N AAC									99 297
				R AGG									119 357
				E GAG									139 417
S TCT	-	D GAC		E GAA									159 47 7
				M ATG								W TGG	179 537
				R AGA									199 597
				Y TAC									219 657
A GCG				S TCC								I ATT	239 717
				Q CAG									259 777

Fig. 4A

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	V GTG																		V GTG	299 897
L CTG	G GGC	W TGG	L CTG	N AAC	L CTG	L CTT	Y TAC	Y TAC	T ACA	R CGG	G GGC	F TTT	Q CAG	H CAC	T ACA			Y TAC		319 957
V GTC	M ATG	I ATC	Q CAG	K AAG	V GTC	I ATC	L CTT	R CGA	D GAC	L CTG	L CTC	R CGT	F TTC	L CTG	L CTG	V GTC	Y TAC	L CTG	V GTC	339 1017
F TTC	L CTT	F TTC	G GGC	F TTT	A GCT	V GTA	A GCC	L CTA	V GTA	S AGC	L TTG	S AGC	R AGA	E GAG	A GCC	R CGA	S AGT	P CCC	K AAA	359 1077
A GCC	P CCT	E GAA		N AAC	N AAC	S TCC	T ACA	V GTG	T ACG	E GAA	Q CAG	P CCC	T ACG	V GTG	G GGC	Q CAG	E GAG	E GAG	E GAG	379 1137
	A GCT																			399 1197
	G GGG																	L TTG		419 1257
L TTG	A GCC		V GTC		L CTC											A GCT				439 1317
TTG E		TAC V	gtc n	CTT H	CTC V	ACC A	TAC D	GTC N	CTG S	CTG W	CTC S	AAC I	ATG W	CTC K	ATT L	GCT Q	CTC K	ATG A	AGC	
TTG E GAA S	GCC T	TAC V GTC L	GTC N AAC E	CTT H CAC M	CTC V GTT E	ACC A GCT N	TAC D GAC G	GTC N AAC Y	S AGC W	CTG W TGG W	S AGC	AAC I ATC R	ATG W TGG R	CTC K AAG K	ATT L TTG K	GCT Q CAG H	CTC K AAA R	ATG A GCC E	AGC I ATC G	1317 459
TTG E GAA S TCT	GCC T ACT V	V GTC L TTG	GTC N AAC E GAG K	CTT H CAC M ATG	V GTT E GAG	ACC A GCT N AAT	D GAC GGT	GTC N AAC Y TAC G	CTG S AGC W TGG D	CTG W TGG W TGG G	S AGC C TGC	AAC I ATC R CGG	W TGG R AGG	CTC K AAG K AAG	ATT L TTG K AAA	GCT Q CAG H CAT	CTC K AAA R CGT C	ATG GCC E GAA	AGC I ATC G GGG R	1317 459 1377 479
E GAA S TCT R AGG	GCC T ACT V GTC L CTG	V GTC L TTG CTG	GTC N AAC E GAG K AAA V	CTT H CAC M ATG V GTC	V GTT E GAG GGC W	ACC A GCT N AAT T ACC	D GAC GGT R AGG	GTC N AAC Y TAC G GGG	S AGC W TGG D GAT	CTG W TGG W TGG GGT K	S AGC C TGC T ACC	AAC I ATC R CGG P CCT	W TGG R AGG D GAT	CTC K AAG K AAG E GAG	L TTG K AAA R CGC	GCT Q CAG H CAT W TGG	CTC K AAA R CGT C TGC	ATG GCC E GAA F TTC	AGC I ATC G GGG R AGG	1317 459 1377 479 1437 499
E GAA S TCT R AGG V GTG S	GCC T ACT V GTC L CTG	V GTC L TTG E GAA	GTC NAAC EGAG KAAA VGTA G	H CAC M ATG V GTC N AAT	V GTT E GAG GGC W TGG	ACC A GCT N AAT T ACC A GCT	D GAC GGT R AGG	GTC NAAC YTAC GGGG WTGG	S AGC W TGG D GAT E GAG	CTG W TGG GGT K AAG	S AGC C TGC ACC	AAC I ATC R CGG P CCT L CTT	W TGG R AGG D GAT P CCC	CTC K AAG K AAG E GAG T ACC	L TTG K AAA R CGC L TTA	Q CAG H CAT W TGG	CTC K AAA R CGT C TGC E GAG	ATG A GCC E GAA F TTC D GAT	AGC I ATC G GGG R AGG P CCA	1317 459 1377 479 1437 499 1497 519

 ${\tt TGGCCCAGATGCAGCAGCAGGCTGGCAGGATGGAGTAGGGAATCTTCCCAGCCACACCAGAGGCTACTGAATTTTGGTG}$

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U.S. Serial No.: 09/587,111
                                          Filed: June 2, 2000
             Title: NOVEL MEMBERS OF THE CAPSAICIN/VANILLOID
             RECEPTOR FAMILY OF PROTEINS AND USES THEREOF
             Agent: Mario Cloutier
              Docket No.: MPI98-093P2RCP3DV1AM
                                  12/37
GAP of: humanvr2.pep check: 5746 from: 1 to: 764
humanVR2 Flh21e11
to: humanvrl.pep check: 6877 from: 1 to: 839
humanVR1 _Fbh18547pat - fchrb87a6, 3909 bases, 4554 checksum.
Symbol comparison table:
/ddm_local/gcg/gcg_9.1/gcgcore/data/rundata/blosum62.cmp
CompCheck: 6430
       Gap Weight:
                     12 Average Match:
                                         2.912
     Length Weight:
                      4 Average Mismatch: -2,003
          Quality: 1530
                                 Length:
                                           850
            Ratio: 2.003
                                   Gaps:
                                            10
Percent Similarity: 55.378 Percent Identity: 46.348
      Match display thresholds for the alignment(s):
                 = IDENTITY
                 : = 2
                  \cdot = 1
humanvr2.pep x humanvr1.pep
                                                 MTSPSSSPVF 10
     1 MKKWSSTDLGTAADPLQKDTCPDPLDGDPNSRPPPAKPQLPTAKSRTRLF 50
    11 RLETLDGGQEDGSEADRGKLDFGSGLPPMESQFQGEDRKFAPQIRVNLNY 60
                       : |.||
    51 GKGDSEEAFPVDCPHEEGELDSCPTI.TVSPVITIQRPGDGPTGARLLSQ 99
    61 RKGTGASQPDPNRFDRDRLFNAVSRGVPEDLAGLPEYLSKTSKYLTDSEY 110
                     :11 :1 11...
                                     :|| | :| |. |:|||.|:
   100 DSVAASTEKTLRLYDRRSIFEAVAONNCODLESLLLFLOKSKKHLTDNEF 149
   111 TEGSTGKTCLMKAVLNLKDGVNACILPLLQIDRDSGNPQPLVNAQCTDDY 160
                                    150 KDPETGKTCLLKAMLNLHDGQNTTIPLLLEIARQTDSLKELVNASYTDSY 199
   161 YRGHSALHIAIEKRSLOCVKLLVENGANVHARACGRFFOKGOG.TCFYFG 209
```

Fig. 5A

200 YKGQTALHIAIERRNMALVTLLVENGADVQAAAHGDFFKKTKGRPGFYFG 249

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Agent: Mario Cloutier

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210	ELPLSLAACTKOWDVVSYLLENPHOPASLQATDSQGNTVLHALVMISDNS	259
250		299
260		309
300	: .: : :: . :	349
310	FRHILOREFSGLSHLSRKFTEWCYGPVRVSLYDLASVDSCEENSVLEI :	357
350	LAYILQREIQEPECRHLSRKFTEWAYGPVHSSLYDLSCIDTCEKNSVLEV	399
	IAF.HCKSPHRHRMVVLEPLNKLLQAKWDLLIPK.FFLNFLCNLIYMFIF	405
	IAYSSSETPNRHDMLLVEPLNRLLQDKWDRFVKRIFYFNFLVYCLYMIIF	449
	TAVAYHQPTLKKQAAPHLKAEVGNSMLLTGHILILLGGIYLLVGQLWYFW :. .: . : :	
	TMAÄYYRPVDGLPPFKMEKIĞDYFRVTĞEİLSVLĞĞVYFFFRGIQYFL	
	RRHVFIWISFIDSYFEILFLFQALLTVVSQVLCFLAIEWYLPLLVSALVL	
	QRRPSMKTLFVDSYSEMLFFLQSLFMLATVVLYFSHLKEYVASMVFSLAL	
	GWLNLLYYTRGFQHTGIYSVMIQKVILRDLLRFLLIYLVFLFGFAVALVS :	
	LSQEAWRPEAPTGPNATESVQPMEGQEDEGNGAQYRGILEASLELFKFTI	
598	:: :	643
	GMGELAFQEQLHFRGMVLLLLLAYVLLTYILLLNMLIALMSETVNSVATD	655
	: : . :: :	693
656	SWSIWKLQKAISVLEMENGYWWC.RKKQRAGVMLTVGTKPDGSPDERWCF	704
694	. : .: : : . :	743
705	RVEEVNWASWEQTLPTLCEDPSGA.GVPRTLENPVLASPPKEDEDGASEE	753
	: · · : · · · RVDEVNWTTWNTNVGIINEDPGNCEGVKRTLSFSLRSSRVSGRHWK	789
754	NYVPVQLLQSN	764
790	NFALVPLLREASARDRQSAQPEEVYLRQFSGSLKPEDAEVFKSPAASGEK	839

Applicant: Rory A. J. Curtis U.S. Serial No.: 09/587,111 **Filed**: June 2, 2000 Title: NOVEL MEMBERS OF THE CAPSAICIN/VANILLOID RECEPTOR FAMILY OF PROTEINS AND USES THEREOF Agent: Mario Cloutier Docket No.: MPI98-093P2RCP3DV1AM 14/37 GAP of: humanvr2.seq check: 8853 from: 1 to: 2809 humanVR2 21e11a, 2809 bases, 8853 checksum. to: humanvrl.seq check: 4554 from: 1 to: 3909 humanVR1 Fbh18547pat - Import - complete Symbol comparison table: /ddm_local/gcg/gcg_9.1/gcgcore/data/rundata/nwsgapdna.cmp CompCheck: 8760 Gap Weight: 50 Average Match: 10.000 Length Weight: Average Mismatch: 0.000 3 Ouality: 14359 3934 Length: Ratio: 5.112 Gaps: 15 Percent Similarity: 55.316 Percent Identity: 55.316 Match display thresholds for the alignment(s): = IDENTITY 5 humanvr2.seq x humanvr1.seq GGCTAGCCTGTCCTGACAGGGGAGAG 26 801 TGTCCACAGTAGTCCCCCCTTATCCACGGGTGTCACTTTCCATGGGTTCA 850 27 TTAAGCTCCCGTTCTCCACCGTGCCGGCTGGCCAGGTGGGCTGAGGGTGA 76 851 GTTATTTGCGGTCAACCACGGTCTGCCAATATTAAATGGAAAATTCTTCA 900 77 CCGAGAGACCAGAACCTGCTTGCTGGAGCTTAGTGCTCAGAGCTGGGGAG 126 901 AACAGTTCCCAAGTTTTCCCTTGTGCATTGTTCTGAGCAGTGTGATGAAG 950 127 GGAGGTTCCGCCGCTCTCTGCTGTCAGCGCCGGCAGCCCCTCCCGGCTT 176

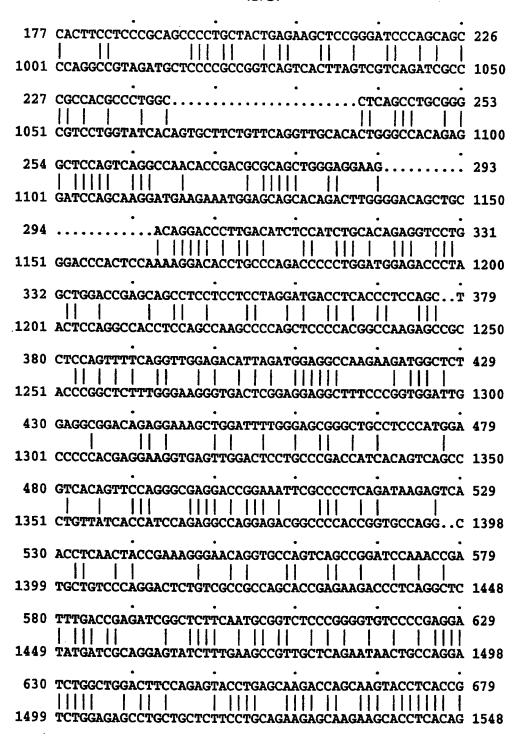
Fig. 6A

951 AGTCTCTGCCGTGCCATCTGGGATGCAAACCGTCCCTGTGTCCCCCACGT 1000

U.S. Serial No.: 09/587,111 Filed: June 2, 2000 Title: NOVEL MEMBERS OF THE CAPSAICIN/VANILLOID RECEPTOR FAMILY OF PROTEINS AND USES THEREOF

Agent: Mario Cloutier

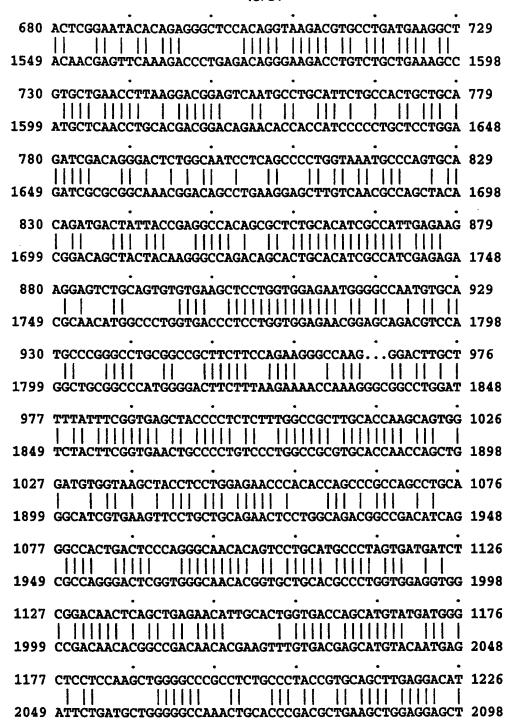
Docket No.: MPI98-093P2RCP3DV1AM



U.S. Serial No.: 09/587,111 Filed: June 2, 2000 Title: NOVEL MEMBERS OF THE CAPSAICIN/VANILLOID RECEPTOR FAMILY OF PROTEINS AND USES THEREOF

Agent: Mario Cloutier

Docket No.: MPI98-093P2RCP3DV1AM



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Agent: Mario Cloutier

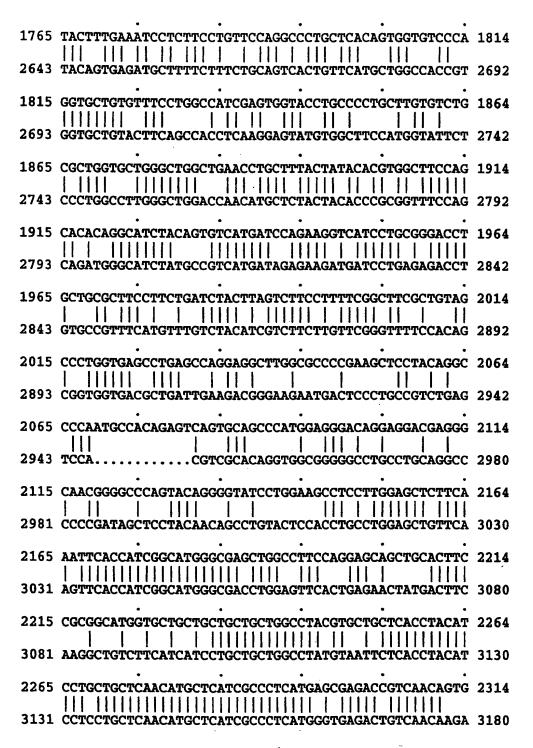
Docket No.: MP198-093P2RCP3DV1AM

1227	CCGCAACCTGCAGGATCTCACGCCTCTGAAGCTGGCCGCCAAGGAGGGCA	1276
2099	CACCAACAAGAAGGGAATGACGCCGCTGGCTCTGGCAGCTGGGACCGGGA	2148
1277	AGATCGAGATTTTCAGGCACATCCTGCAGCGGGAGTTTTCAGGA	1320
2149	AGATCGGGGTCTTGGCCTATATTCTCCAGCGGGAGATCCAGGAGCCCGAG	2198
1321	CTGAGCCACCTTTCCCGAAAGTTCACCGAGTGGTGCTATGGGCCTGTCCG	1370
2199	TGCAGGCACCTGTCCAGGAAGTTCACCGAGTGGGCCTACGGGCCCGTGCA	2248
1371	GGTGTCGCTGTATGACCTGGCTTCTGTGGACAGCTGTGAGGAGAACTCAG	1420
2249	CTCCTCGCTGTACGACCTGTCCTGCATCGACACCTGCGAGAAGAACTCGG	2298
1421	TGCTGGAGATCATTGCCTTTCATTGCAAGAGCCCGCACCGACACCGA	1467
2299	TGCTGGAGGTGATCGCCTACAGCAGCAGCGAGACCCCTAATCGCCACGAC	2348
1468	ATGGTCGTTTTGGAGCCCCTGAACAAACTGCTGCAGGCGAAATGGGA	1514
2349	ATGCTCTTGGTGGAGCCGCTGAACCGACTCCTGCAGGACAAGTGGGACAG	2398
1515	TCTGCTCATCCCCAAGTTCTTCTTAAACTTCCTGTGTAATCTGATCTACA	1564
2399	ATTCGTCAAGCGCATCTTCTACTTCAACTTCCTGGTCTACTGCCTGTACA	2448
1565	TGTTCATCTTCACCGCTGTTGCCTACCATCAGCCTACCCTGAAGAAGCAG	1614
2449	TGATCATCTTCACCATGGCTGCCTACTACAGGCCCGTGGATGGCTT	2494
1615	GCCGCCCTCACCTGAAAGCGGAGGTTGGAAACTCCATGCTGCTGACGGG	1664
2495	ĠĊĊŦĊĊĊŦŤŦĂAĠĂŦĠĠAAAÄAAŤŤĠĠĀĠĀĊŤAŦŦŤĊĊĠAĠŤŦĀĊŦĠĠ	2542
1665	CCACATCCTTATCCTGCTAGGGGGGATCTACCTCCTCGTGGGCCAGCTGT	1714
2543	AGAGATCCTGTCTGTTAGGAGGAGTCTACTTCTTTTTCCGAGGGATTC	2592
1715	GGTACTTCTGGCGGCGCCACGTGTTCATCTGGATCTCGTTCATAGACAGC	1764
2593	AGTATTTCCTGCAGAGGCGGCCGTCGATGAAGACCCTGTTTGTGGACAGC	2642

U.S. Serial No.: 09/587,111 Filed: June 2, 2000 Title: NOVEL MEMBERS OF THE CAPSAICIN/VANILLOID RECEPTOR FAMILY OF PROTEINS AND USES THEREOF

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Agent: Mario Cloutier

Docket No.: MPI98-093P2RCP3DV1AM

2315	${\tt TCGCCACTGACAGCTGGAGCATCTGGAAGCTGCAGARAGCCATCTCTGTC}$	2364
3181	TCGCACAGGAGACAACATCTGGAAGCTGCAGAGAGCCATCACCATC	3230
2365	CTGGAGATGGAGAATGGCTATTGGTGGTGCAGGAAGAAGCAGCGGGC	2411
3231	CTGGACACGGAGAAGAGCTTCCTTAAGTGCATGAGGAAGGCCTTCCGCTC	3280
2412	AGGTGTGATGCTGACCGTTGGCACTAAGCCAGATGGCAGCCCGGATGAGC	2461
3281	AGGCAAGCTGCAGGTGGGGTACACCCTGATGGCAAGGACGACTACC	3330
2462	GCTGGTGCTTCAGGGTGGAGGAGGTGAACTGGGCTTCATGGGAGCAGACG	2511
	GGTGGTGCTTCAGGGTGGACGAGGTGAACTGGACCACCCAGC	
2512	CTGCCTACGCTGTGAGGACCCGTCAGGGGCAGGTGTCCCTCGAAC	2558
	GTGGGCÀTCATCAACGÀAGÀCCCGGGCAACTGTGAGGGCGTCAAGCGCÀC	
	TCTCGAGAACCCTGTCCTGGCTTCCCCTCCCAAGGAGGATGAGGAT	
	CCTGAGCTTCTCCCTGCGGTCAAGCAGAGTTTCAGGCAGACACTGGAAGA	
	GGTGCCTCTGAGGAAAACTATGTGCCCGTCCAGCTCCCAGTCCAACTG	
	ACTTTGCCCTGGTCCCCCTTTTAAGAGAGGCAAGTGCTCGAGATAGGCAG	
	ATGGCCCAGATGCAGCAGGAGGCCAGAGGACAGAGCAGAGCATCTTTCCA	
	TCTGCTCAGCCCGAGGAAGTTTATCTGCGACAGTTTTCAGGGTCTCTGAA	
	ACCACATCTGCTGGCTCTGGGGTCCCAGTGAATTCTGGTGGCAAATATAT	
	ATTTTCACTAACTCAAAAAAAAAAAAAAAAAAAAAAAA	
	ATTITCACTAACTCAAAAAAAAAAAAAAAAAAAAAAAAA	
	AAAAA	
	CCCCGTTGCCACGGGGGCTGCTGAGGGAACACCAGTGCTCTGTCAGCAG	

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CLUSTAL W (1.74) multiple sequence alignment

humanVR2 rat VR2	MTSPSSSPVFRLETLDGGQEDGSEADRGKLDFGSGLPPMESQFQGEDRKFAPQIRVNLNY
humanVR2 rat VR2	RKGTGASQPDPNRFDRDRLFNAVSRGVPEDLAGLPEYLSKTSKYLTDSEYTEGSTGKTCL
humanVR2 rat VR2	MKAVLNLKDGVNACILPLLQIDRDSGNPQPLVNAQCTDDYYRGHSALHIAIEKRSLQCVK
humanVR2 rat VR2	LLVENGANVHARACGRFFQKGQGTCFYFGELPLSLAACTKQWDVVSYLLENPHQPASLQASTHASALSLAACTKQWDVVTYLLENPHQPASLEA .************************************
humanVR2 rat VR2	TDSQGNTVLHALVMISDNSAENIALVTSMYDGLLQAGARLCPTVQLEDIRNLQDLTPLKL TDSLGNTVLHALVMIADNSPENSALVIHMYDGLLQMGARLCPTVQLEEISNHQGLTPLKL
humanVR2 rat VR2	AAKEGKIEIFRHILQREFSG-LSHLSRKFTEWCYGPVRVSLYDLASVDSCEENSVLEIIA AAKEGKIEIFRHILQREFSGPYQPLSRKFTEWCYGPVRVSLYDLSSVDSWEKNSVLEIIA
humanVR2 rat VR2	FHCKSPHRHRMVVLEPLNKLLQAKWDLLIPKFFLNFLCNLIYMFIFTAVAYHQPTLKKQA FHCKSPNRHRMVVLEPLNKLLQEKWDRLVSRFFFNFACYLVYMFIFTVVAYHQPSLDQPA
humanVR2 rat VR2	APHLKAEVGNSMLLTGHILILLGGIYLLVGQLWYFWRRHVFIWISFIDSYFEILFLFQAL IPSSKATFGESMLLLGHILILLGGIYLLLGQLWYFWRRRLFIWISFMDSYFEILFLLQAL * * .*:*** ****************************
humanVR2 rat VR2	LTVVSQVLCFLAIEWYLPLLVSALVLGWLNLLYYTRGFQHIGIYSVMIQKVILRDLLRFL LTVLSQVLRFMETEWYLPLLVLSLVLGWLNLLYYTRGFQHTGIYSVMIQKVILRDLLRFL ***:*** *: ******** :*****************
humanVR2 rat VR2	LIYLVFLFGFAVALVSLSQEAWRPEAPTGPNATESVQPMEGQEDEGNGAQYRGILEASLE LVYLVFLFGFAVALVSLSREARSPKAPEDNNSTVTEQPTVGQEEEPAPYRSILDASLE *:***********************************
humanVR2 rat VR2	LFKFTIGMGELAFQEQLHFRGMVLLLLLAYVLLTYILLLNMLIALMSETVNSVATDSWSI LFKFTIGMGELAFQEQLRFRGVVLLLLLAYVLLTYVLLLNMLIALMSETVNHVADNSWSI
humanVR2 rat VR2	WKLQKAISVLEMENGYWWCR-KKQRAGVMLTVGTKPDGSPDERWCFRVEEVNWASWEQTL WKLQKAISVLEMENGYWWCRRKKHREGRLLKVGTRGDGTPDERWCFRVEEVNKAAWEKTL
humanVR2 rat VR2	PTLCEDPSGAGVPRTLENPVLASPPKEDEDGASEENYVPVQLLQSN PTLSEDPSGPGITGNKKNPTSK-PGKNSASEEDHLPLQVLQSP ***.*********************************

Applicant: Rory A. J. Curtis U.S. Serial No.: 09/587,111 Filed: June 2, 2000 Title: NOVEL MEMBERS OF THE CAPSAICIN/VANILLOID RECEPTOR FAMILY OF PROTEINS AND USES THEREOF Agent: Mario Cloutier Docket No.: MPI98-093P2RCP3DV1AM 21/37 GAP of: ratvr2.pep check: 9190 from: 1 to: 554 ratVR2 Flrxb147g11 to: humanvr2.pep check: 5746 from: 1 to: 764 humanVR2 Flh21e11 Symbol comparison table: /usr/local/gog_9.1/gcgcore/data/rundata/blosum62.cmp CompCheck: 6430 Gap Weight: 12 Average Match: 2.912 Length Weight: Average Mismatch: -2.003 766 Quality: 2182 Length: Ratio: 3.939 Gaps: Percent Similarity: 81.703 Percent Identity: 79.167 Match display thresholds for the alignment(s): = IDENTITY = 2. = 1 ratvr2.pep x humanvr2.pep 1STHASALSLAACTKOWDVVTYLLENPHOPASLEATDSLGNTVLH 44 201 GQGTCFYFGELPLSLAACTKQWDVVSYLLENPHQPASLQATDSQGNTVLH 250 45 ALVMIADNSPENSALVIHMYDGLLOMGARLCPTVOLEEISNHOGLTPLKL 94 251 ALVMISDNSAENIALVTSMYDGLLQAGARLCPTVQLEDIRNLQDLTPLKL 300 95 AAKEGKIEIFRHILOREFSGPYOPLSRKFTEWCYGPVRVSLYDLSSVDSW 144 301 AAKEGKIEIFRHILQREFSG.LSHLSRKFTEWCYGPVRVSLYDLASVDSC 349 145 EKNSVLEIIAFHCKSPNRHRMVVLEPLNKLLQEKWDRLVSRPFFNFACYL 194

Fig. 8A

350 EENSVLEIIAFHCKSPHRHRMVVLEPLNKLLOAKWDLLIPKFFLNFLCNL 399

195 VYMFIFTVVAYHQPSLDQPAIPSSKATFGESMLLLGHILILLGGIYLLLG 244

400 IYMFIFTAVAYHOPTLKKOAAPHLKAEVGNSMLLTGHILILLGGIYLLVG 449

U.S. Serial No.: 09/587,111 Filed: June 2, 2000 Title: NOVEL MEMBERS OF THE CAPSAICIN/VANILLOID RECEPTOR FAMILY OF PROTEINS AND USES THEREOF

Agent: Mario Cloutier

Docket No.: MPI98-093P2RCP3DV1AM

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245	QLWYFWRRLFIWISFMDSYFEILFLLQALLTVLSQVLRFMETEWYLPLL	294
450	QLWYFWRRHVFIWISFIDSYFEILFLFQALLTVVSQVLCFLAIEWYLPLL	499
	VLSLVLGWLNLLYYTRGFQHTGIYSVMIQKVILRDLLRFLLVYLVFLFGF	
	VSALVLGWLNLLYYTRGFQHTGIYSVMIQKVILRDLLRFLLIYLVFLFGF	
	AVALVSLSREARSPKAPEDNNSTVTEQPTVGQEEEPAPYRSILDASLE	
	LFKFTIGMGELAFQEQLRFRGVVLLLLAYVLLTYVLLLNMLIALMSETV	
	NHVADNSWSIWKLQKAISVLEMENGYWWCRRKKHREGRLLKVGTRGDGTP	
	.	-
493	DERWCFRVEEVNWAAWEKTLPTLSEDPSGPGITGNKKNPTSKPGKN	538
699	DERWCFRVEEVNWASWEQTLPTLCEDPSGAGVPRTLENPVLASPPKEDED	748
539	SASEEDHLPLQVLQSP 554	
749	GASEENYVPVOLLOSN 764	

Fig. 8B

RECEPTOR FAMILY OF PROTEINS AND USES THEREOF Agent: Mario Cloutier Docket No.: MPI98-093P2RCP3DV1AM 23/37 GAP of: humanvr1.seq check: 4554 from: 1 to: 3909 humanVR1 Fbh18547pat - Import - complete to: ratvr1.seq check: 7921 from: 1 to: 2847 ratVR1.seq AF029310 in GenBank Symbol comparison table: /ddm_local/gcg/gcg_9.1/gcgcore/data/rundata/nwsgapdna.cmp CompCheck: 8760 Gap Weight: 50 Average Match: 10.000 Length Weight: 3 Average Mismatch: 0.000 Quality: 22717 Length: 3914 Ratio: 7.979 Gaps: 10 Percent Similarity: 82.125 Percent Identity: 82.125 Match display thresholds for the alignment(s): = IDENTITY : = 5 . = 1 humanvrl . seq x ratvrl . seq 1001 CCAGGCCGTAGATGCTCCCCGCCGGTCAGTCACTTAGTCGTCAGATCGCC 1050 .CAGCTCCAAGGCACTTGCTCC 21 22 ATTTGGGGTGTGCCTGCACCT...AGCTGGTTGCAAATTGGGCCACAGAG 68 1101 GATCCAGCAAGGATGAAGAAATGGAGCACCAGACTTGGGGACAGCTGC 1150

Applicant: Rory A. J. Curtis U.S. Serial No.: 09/587,111

Filed: June 2, 2000

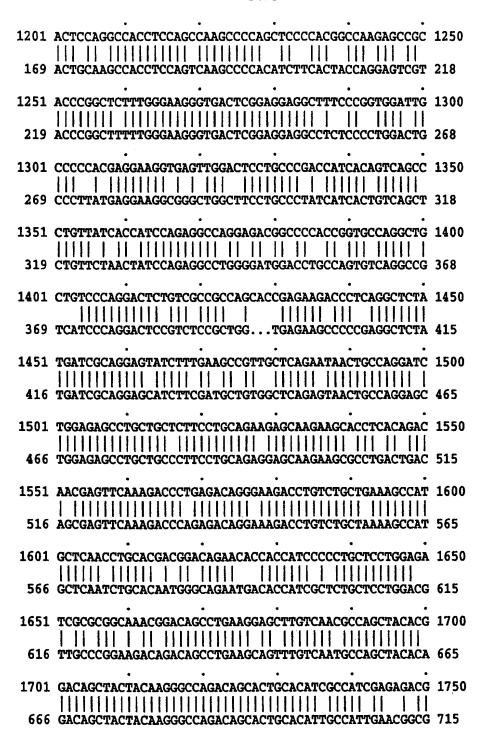
Title: NOVEL MEMBERS OF THE CAPSAICIN/VANILLOID

Fig. 9A

U.S. Serial No.: 09/587,111 Filed: June 2, 2000 Title: NOVEL MEMBERS OF THE CAPSAICIN/VANILLOID RECEPTOR FAMILY OF PROTEINS AND USES THEREOF

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Agent: Mario Cloutier

Docket No.: MPI98-093P2RCP3DV1AM

1751	CAACATGCCCTGGTGACCCTCCTGGTGGAGACGGAGCAGACGTCCAGG	1800
716	GAACATGACGCTGGTGACCCTCTTGGTGGAGAATGGAGCAGATGTCCAGG	765
1801	CTGCGGCCCATGGGGACTTCTTTAAGAAAACCAAAGGGCGGCCTGGATTC	1850
766	CTGCGGCTAACGGGACTTCTTCAAGAAAACCAAAGGGAGGCCTGGCTTC	815
1851	TACTTCGGTGAACTGCCCCTGTCCCTGGCCGCGTGCACCAACCA	1900
816	TACTTTGGTGAGCTGCCCCTGTCCCTGGCTGCGTGCACCAACCA	865
1901	CATCGTGAAGTTCCTGCTGCAGAACTCCTGGCAGACGGCCGACATCAGCG	1950
866	CATTGTGAAGTTCCTGCAGAACTCCTGGCAGCCTGCAGACATCAGCG	915
1951	CCAGGGACTCGGTGGCAACACGGTGCTGCACGCCCTGGTGGAGGTGGCC	2000
	CCCGGGACTCAGTGGGCAACACGGTGCTTCATGCCCTGGTGGAGGTGGCA	
	GACAACACGGCCGACAACACGAAGTTTGTGACGAGCATGTACAATGAGAT	
	GATAACACAGTTGACAACACCAAGTTCGTGACAAGCATGTACAACGAGAT	
	TCTGATGCTGGGGGCCAAACTGCACCCGACGCTGAAGCTGGAGGAGCTCA	
	CTTGATCCTGGGGGCCAAACTCCACCCCACGCTGAAGCTGGAAGAGATCA	
	CCAACAAGAAGGGAATGACGCCGCTGGCTCTGGCAGCTGGGACCGGGAAG	
	ATCGGGGTCTTGGCCTATATTCTCCAGCGGGAGATCCAGGAGCCCGAGTG	
	ATCGGGGTCTTGGCCTACATTCTCCAGAGGGAGATCCATGAACCCGAGTG	
	CAGGCACCTGTCCAGGAAGTTCACCGAGTGGGCCTACGGGCCCGTGCACT	
	CCTCGCTGTACGACCTGTCCTGCATCGACACCTGCGAGAAGAACTCGGTG	

Applicant: Rory A. J. Curtis
U.S. Serial No.: 09/587,111 Filed: June 2, 2000
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Agent: Mario Cloutier

Docket No.: MPI98-093P2RCP3DV1AM

2301	CTGGAGGTGATCGCCTACAGCAGCAGCGAGACCCCTAATCGCCACGACAT	2350
1266	CTGGAGGTGATCGCTTACAGCAGCAGTGAGACCCCTAACCGTCATGACAT	1315
2351	GCTCTTGGTGGAGCCGCTGAACCGACTCCTGCAGGACAAGTGGGACAGAT	2400
1316	GCTTCTCGTGGAACCCTTGAACCGACTCCTACAGGACAAGTGGGACAGAT	1365
2401	TCGTCAAGCGCATCTTCTACTTCAACTTCCTGGTCTACTGCCTGTACATG	2450
1366		1415
2451	ATCATCTTCACCATGGCTGCCTACTACAGGCCCGTGGATGGCTTGCCTCC	2500
	ATCATCTTCACCGCGGCTGCCTACTATCGGCCTGTGGAAGGCTTGCCCCC	
	CTTTAAGATGGAAAAATTGGAGACTATTTCCGAGTTACTGGAGAGA	
	CTATAAGCTGAAAAACACCGTTGGGGACTATTTCCGAGTCACCGGAGAGA	
	TCCTGTCTGTGTTAGGAGGAGTCTACTTCTTTTTCCGAGGGATTCAGTAT	
1516 2598		1565
	TTCCTGCAGAGGCGACCATCCCTCAAGAGTTTGTTGTGGACAGCTACAG	
2648		2697
2698		2747
1666	TGTACTTCAGCCAACGCAAGGAGTATGTGGCTTCCATGGTGTTCTCCCTG	1715
2748	GCCTTGGGCTGGACCAACATGCTCTACTACACCCGCGGTTTCCAGCAGAT	2797
1716	GCCATGGGCTGGACCAACATGCTCTACTATACCCGAGGATTCCAGCAGAT	1765
2798	GGGCATCTATGCCGTCATGATAGAGAAGATGATCCTGAGAGACCTGTGCC	2847
1766	GGGCATCTATGCTGTCATGATTGAGAAGATGATCCTCAGAGACCTGTGCC	1815

Fig. 9D

Filed: June 2, 2000 Title: NOVEL MEMBERS OF THE CAPSAICIN/VANILLOID RECEPTOR FAMILY OF PROTEINS AND USES THEREOF

Agent: Mario Cloutier

Docket No.: MPI98-093P2RCP3DV1AM

2848	GTTTCATGTTTGTCTACATCGTCTTCTTGTTCGGGTTTTCCACAGCGGTG	2897
1816	GGTTTATGTTCGTCTACCTCGTGTTCTTGTTTGGATTTTCCACAGCTGTG	1865
2898	GTGACGCTGATTGAAGACGGGAAGAATGACTCCCTGCCGTCTGAGTCCAC	2947
1866	GTGACACTGATTGAGGATGGGAAGAATAACTCTCTGCCTATGGAGTCCAC	1915
2948	GTCGCACAGGTGGCGGGGCCTGCCTGCAGGCCCCCGATAGCTCCTACA	2997
1916	ACCACACAAGTGCCGGGGTCTGCCTGCAAGCCAGGTAACTCTTACA	1962
2998	ACAGCCTGTACTCCACCTGCCTGGAGCTGTTCAAGTTCACCATCGGCATG	3047
1963	ACAGCCTGTATTCCACATGTCTGGAGCTGTTCAAGTTCACCATCGGCATG	2012
3048	GGCGACCTGGAGTTCACTGAGAACTATGACTTCAAGGCTGTCTTCATCAT	3097
2013	GGCGACCTGGAGTTCACTGAGAACTACGACTTCAAGGCTGTCTTCATCAT	2062
3098	CCTGCTGCCTGTAATTCTCACCTACATCCTCCTGCTCAACATGC	3147
	CCTGTTACTGGCCTATGTGATTCTCACCTACATCCTTCTGCTCAACATGC	
3148	TCATCGCCCTCATGGGTGAGACTGTCAACAAGATCGCACAGGAGGAGCAAG	3197
	TCATTGCTCTCATGGGTGAGACCGTCAACAAGATTGCACAAGAGAGCAAG	
	AACATCTGGAAGCTGCAGAGAGCCATCACCATCCTGGACACGGAGAAGAG	
	AACATCTGGAAGCTGCAGAGAGAGCCATCCTGGATACAGAGAAGAG	
	CTTCCTTAAGTGCATGAGGAAGGCCTTCCGCTCAGGCAAGCTGCTGCAGG	
	CTTCCTGAAGTGCATGAGGAAGCCTCCGCTCTGGCAAGCTGCTGCAGG	
3298	TGGGGTACACCTGATGGCAAGGACGACTACCGGTGGTGCTTCAGGGTG	3347
	GACGAGGTGAACTGGACCACCTGGAACACCAACGTGGGCATCATCAACGA	
2313	GACGAGGTAAACTGGACTACCTGGAACACCAATGTGGGTATCATCAACGA	2362

Fig. 9E

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Agent: Mario Cloutier

Docket No.: MPI98-093P2RCP3DV1AM

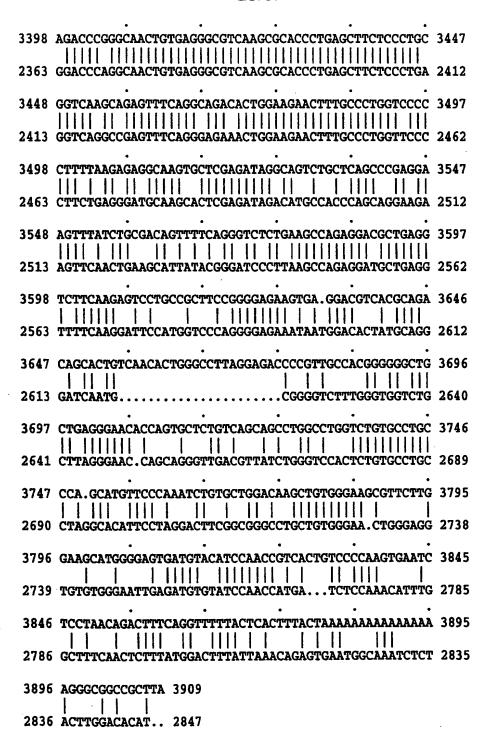


Fig. 9F

Title: NOVEL MEMBERS OF THE CAPSAICIN/VANILLOID RECEPTOR FAMILY OF PROTEINS AND USES THEREOF Agent: Mario Cloutier Docket No.: MPI98-093P2RCP3DV1AM 29/37 GAP of: humanvrl.pep check: 6877 from: 1 to: 839 humanVR1 Fbh18547pat - fchrb87a6, 3909 bases, 4554 checksum. to: ratvr1.pep check: 5764 from: 1 to: 838 ratVR1 | AF029310 Rattus norvegicus vanilloid receptor subtype 1 mRNA, complete cds. Symbol comparison table: /ddm_local/gcg/gcg_9.1/gcgcore/data/rundata/blosum62.cmp CompCheck: 6430 Gap Weight: 12 Average Match: 2.912 Length Weight: Average Mismatch: -2.003 Length: 840 Quality: Ratio: 4.456 Gaps: Percent Identity: 86.022 Percent Similarity: 89.247 Match display thresholds for the alignment(s): = IDENTITY humanvr1.pep x ratvr1.pep 1 MKKWSSTDLGTAADPLQKDTCPDPLDGDPNSRPPPAKPQLPTAKSRTRLF 50 . | |...|.||.|.|||.:||| || : | :|||||| 1 MEORASLDSEESESPPOENSCLDPPDRDPNCKPPPVKPHIFTTRSRTRLF 50 51 GKGDSERAFPVDCPHEEGELDSCPTITVSPVITIQRPGDGPTGARLLSQD 100 51 GKGDSEEASPLDCPYEEGGLASCPIITVSSVLTIQRPGDGPASVRPSSQD 100 101 SVAASTEKTLRLYDRRSIFEAVAQNNCQDLESLLLFLQKSKKHLTDNEFK 150 101 SVSAG.EKPPRLYDRRSIFDAVAQSNCQELESLLPFLQRSKKRLTDSEFK 149 151 DPETGKTCLLKAMLNIHDGQNTTIPLLLEIARQTDSLKELVNASYTDSYY 200 150 DPETGKTCLLKAMLNLHNGQNDTIALLLLDVARKTDSLKQFVNASYTDSYY 199 201 KGQTALHIAIERRNMALVTLLVENGADVQAAAHGDFFKKTKGRPGFYFGE 250 200 KGQTALHIAIERRMTLVTLLVENGADVQAAANGDFFKKTKGRPGFYFGE 249

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Fig. 10A

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251	LPLSLAACTNQLGIVKFLLQNSWQTADISARDSVGNTVLHALVEVADNTA 300
250	
301	DNTKFVTSMYNEILMLGAKLHPTLKLEELTNKKGMTPLALAAGTGKIGVL 350
300	DNTKFVTSMYNEILILGAKLHPTLKLEEITNRKGLTPLALAASSGKIGVL 349
351	AYILOREIOEPECRHLSRKFTEWAYGPVHSSLYDLSCIDTCEKNSVLEVI 400
350	AYILQREIHEPECRHLSRKFTEWAYGPVHSSLYDLSCIDTCEKNSVLEVI 399
	AYSSSETPNRHDMLLVEPLNRLLQDKWDRFVKRIFYFNFLVYCLYMIIFT 450
	AYSSSETPNRHDMLLVEPLNRLLQDKWDRFVKRIFYFNFFVYCLYMIIFT 449
	MAAYYRPVDGLPPFKMEK.IGDYFRVTGEILSVLGGVYFFFRGIQYFLQR 499
	AAAYYRPVEGLPPYKLKNTVGDYFRVTGEILSVSGGVYFFFRGIQYFLQR 499 RPSMKTLFVDSYSEMLFFLQSLFMLATVVLYFSHLKEYVASMVFSLALGW 549
•	: . : RPSLKSLFVDSYSEILFFVQSLFMLVSVVLYFSQRKEYVASMVFSLAMGW 549
550	
550	
600	EDGKNDSLPSESTSHRWRGPACRPPDSSYNSLYSTCLELFKFTIGMGDLE 649
600	. . : :
650	FTENYDFKAVFIILLLAYVILTYILLLNMLIALMGETVNKIAQESKNIWK 699
649	FTENYDFKAVFIILLLAYVILTYILLINMLIALMGETVNKIAQESKNIWK 698
	LQRAITILDTEKSFLKCMRKAFRSGKLLQVGYTPDGKDDYRWCFRVDEVN 749
	LQRAITILDTEKSFLKCMRKAFRSGKLLQVGFTPDGKDDYRWCFRVDEVN 748
	WTTWNTNVGIINEDPGNCEGVKRTLSFSLRSSRVSGRHWKNFALVPLLRE 799
	WITWITHVGIINEDPGNCEGVKRTLSFSLRSGRVSGRNWKNFALVPLLRD 798
	ASARDRQSAQPEEVYLRQFSGSLKPEDAEVFKSPAASGEK 839 . : :. ASTRDRHATQQEEVQLKHYTGSLKPEDAEVFKDSMVPGEK 838
133	PATIMIMUTÄÄRRAÄMMITTÄRHÜL KRUEALINDMATOMI. 000

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CLUSTAL W (1.74) multiple sequence alignment

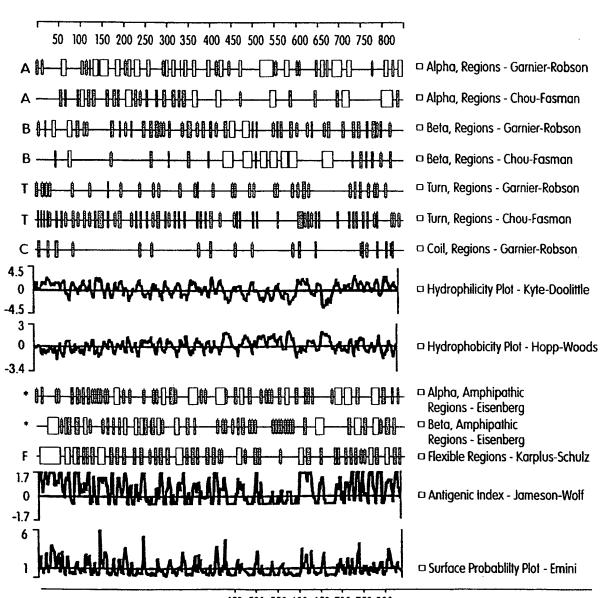
humanVR2.alt human VR2	MTSPSSSPVFRLETLDGGQEDGSEADRGKLDFGSGLPPMESQFQGEDRKFAPQIRVNLNY
humanVR2.alt human VR2	RKGTGASQPDPNRFDRDRLFNAVSRGVPEDLAGLPEYLSKTSKYLTDSEYTEGSTGKTCL
humanVR2.alt human VR2	MKAVLNLKDGVNACILPLLQIDRDSGNPQPLVNAQCTDDYYRGHSALHIAIEKRSLQCVK
humanVR2.alt human VR2	GRFFQKGQGTCFYFGELPLSLAACTKQWDVVSYLLENPHQPASLQA LLVENGANVHARACGRFFQKGQGTCFYFGELPLSLAACTKQWDVVSYLLENPHQPASLQA
humanVR2.alt human VR2	TDSQGNTVLHALVMISDNSAENIALVTSMYDGLLQAGARLCPTVQLEDIRNLQDLTPLKL TDSQGNTVLHALVMISDNSAENIALVTSMYDGLLQAGARLCPTVQLEDIRNLQDLTPLKL
humanVR2.alt human VR2	AAKEGKIEIFRHILQREFSGLSHLSRKFTEWCYGPVRVSLYDLASVDSCEENSVLEIIAF AAKEGKIEIFRHILQREFSGLSHLSRKFTEWCYGPVRVSLYDLASVDSCEENSVLEIIAF
humanVR2.alt human VR2	HCKSPHRHRMVVLEPLNKLLQAKWDLLIPKFFDNFLCNLIYMFIFTAVAYHQPTLKKQAA HCKSPHRHRMVVLEPLNKLLQAKWDLLIPKFFLNFLCNLIYMFIFTAVAYHQPTLKKQAA
humanVR2.alt human VR2	PHLKAEVGNSMLLTGHILILLGGIYLLVGQLWYFWRRHVFIWISFIDSYFEILFLFQALL PHLKAEVGNSMLLTGHYLILLGGIYLLVGQLWYFWRRHVFIWISFIDSYFEILFLFQALL
humanVR2.alt human VR2	TVVSQVLCFLAIEWYLPLLVSALVLGWLNLLYYTRGFQHTGIYSVMIQTVVSQVLCFLAIEWYLPLLVSALVLGWLNLLYYTRGFQHTGIYSVMIQKVILRDLLRFLL
humanVR2.alt human VR2	IYLVFLFGFAVALVSLSQEAWRPEAPTGPNATESVQPMEGQEDEGNGAQYRGILEASLEL
humanVR2.alt human VR2	FKFTIGMGELAFQEQLHFRGMVLLLLLAYVLLTYILLLNMLIALMSETVNSVATDSWSIW
humanVR2.alt human VR2	KKAISVLEMENGYWWCRKKQRAGVMLTVGTKPDGSPDERWCFRVEEVNWASWEQTLPT KLQKAISVLEMENGYWWCRKKQRAGVMLTVGTKPDGSPDERWCFRVEEVNWASWEQTLPT :************************************
humanVR2.alt human VR2	LCEDPSGAGVPRTLENPVLASPPKEDEDGASEENYVPVQLLQSN LCEDPSGAGVPRTLENPVLASPPKEDEDGASEENYVPVQLLQSN ************************************

Fig. 11

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450 500 550 600 650 700 750 800

Fig. 12

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Protein Family / Domain Matches, HMMer version 2

Searching for complete domains hmmpfam - search a single seg against HMM database HMMER 2.1.1 (Dec 1998) Copyright (C) 1992-1998 Washington University School of Medicine HMMER is freely distributed under the GNU General Public License (GPL). /prod/ddm/seganal/PFAM/pfam4.2/Pfam HMM file: /usr/ns-home/docs/seganal/orfanal/oa-script 18670 seg Sequence file: Query: hVR-1 Scores for sequence family classification (score includes all domains): Model Description Score E-value 51.5 1.9e-11 3 ank Ank repeat Parsed for domains: score E-value Model Domain seg-f seg-t hmm-f hmm-t 1/3 201 233 .. 1 33 [] 34.4 2.6e-06 ank 283 .. 1 13.2 2 248 33 [] ank 2/3 26 361 ... 1 33 [] 3.4 ank 3/3 333 Alignments of top-scoring domains: ank: domain 1 of 3, from 201 to 233: score 34.4, E = 2.6e-06 *->nGnTPLHlAarygnvevvklLLehGAdvnartk<-* +G+T+LH+A + n+ +v lL+e+GAdv a+ 233 KGOTALHIAIERRNMALVTLLVENGADVQAAAH hVR-1 201 ank: domain 2 of 3, from 248 to 283: score 13.2, E = 2 *->nGnTPLHlAarygnvevvklLLe...hGAdvnartk<-* G PL 1Aa ++++ +vk+LL+++ + Ad+ ar+ FGELPLSLAACTNQLGIVKFLLQnswQTADISARDS 283 hVR-1 248 domain 3 of 3, from 333 to 361: score 3.4, E = 26 *->nGnTPLHlAarygnvevvklLLehGAdvnartk<-* +G TPL lAa +g++ v ++ L+

Fig. 13

361

KGMTPLALAAGTGKIGVLAYILQ----REIQEP

hvr-1

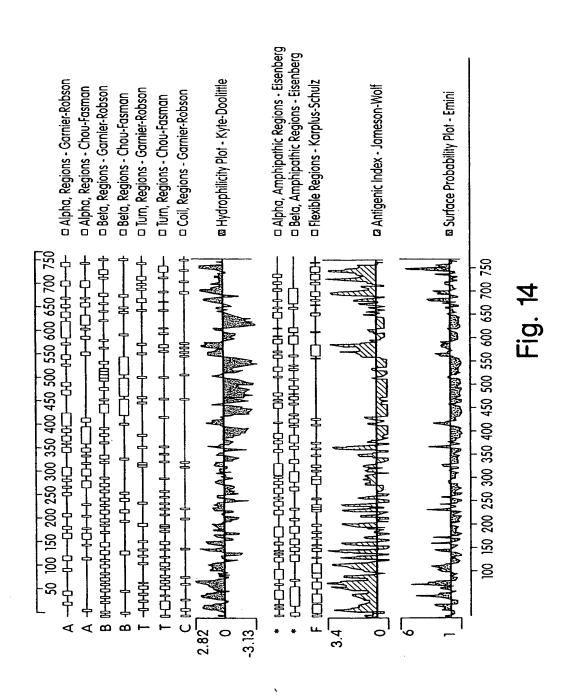
333

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35/37 Protein Family / Domain Matches, HMMer version 2 Searching for complete domains hmmpfam - search a single seq against HMM database HMMER 2.1.1 (Dec 1998) Copyright (C) 1992-1998 Washington University School of Medicine HIMMER is freely distributed under the GNU General Public License (GPL). HMM file: /prod/ddm/seqanal/PFAM/pfam4.2/Pfam Sequence file: /tmp/orfanal.5/g.aa Query: Flh21e11 Scores for sequence family classification (score includes all domains): Model Description Score E-value N ank PF00023 Ank repeat 53.7 4e-12 Parsed for domains: Model Domain seq-f seq-t hmm-f hmm-t score E-value ank 194 .. 1/3 162 1 33 [] 38.3 1.7e-07 2/3 ank 208 243 ... 1 33 [] 6.4 4.3 ank 3/3 293 328 ... 1 33 [] 8.8 2.1 Alignments of top-scoring domains: ank: domain 1 of 3, from 162 to 194: score 38.3, E = 1.7e-07 *->nGnTPLHlAarygnvevvklLLehGAdvnartk<-* +G+++LH+A ++ ++++vklL+e+GA+v+ar Flh21e11 162 RGHSALHIAIEKRSLQCVKLLVENGANVHARAC 194 ank: domain 2 of 3, from 208 to 243: score 6.4, E = 4.3*->nGnTPLHlAarygnvavvklLLe...hGAdvnartk<-* G PL lAa + +++vv +LLe++++ A+ a++ Flh21e11 208 **FGELPLSLAACTKQWDVVSYGLEnphQPASLQATDS** 243 domain 3 of 3, from 293 to 328: score 8.8, E = 2.1 *->nGnTPLHLAarygnvevvklLLe...hGAdvnart<<-* + +TPL lAa++g++e+ + L+++ G + +r Flh21e11 293 **QDLTPLKLAAKEQKLEIFRHILQrafSGLSHLSRK** 328

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Fig. 15

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llvenganvharacgrffokgogtcfyfgelplslaactkowdvvsyllenphopasloatsogntvlhalvm **RKGTGASQPDPNRFDRDRLFNAVSRGVPEDLAGLPEYLSKTSKYLTDSEYTEGSTGKTCL** I SDNSAENI ALVTSMYDGLLQAGARLCPTVQLEDIRNLQDLTPLKLAAKEGKIEIFRHIL **QREFSGLSHLSRKFTEWCYGPVRVSLYDLASVDSCEENSVLEIIAFHCKSPHRHRMVVLE** PLNKLLQAKWDLLIPKFFLNFLCNLIYMFIFTAVAYHQPTLKKQAAPHLKAEVGNSMLLT Ylplivsalvlgwinilyytrgfohtgiysvmiokkaisvlemengywwcrkkoragvml **TVGTKPDGSPDERWCFRVEEVNWASWEQTLPTLCEDPSGAGVPRTLENPVLASPPKEDED** mtspssspvfrletldgggedgseadrgkldfgglppmesqfqgedrkfapqirvnlny MKAVLNIKDGVNACILPLLQIDRDSGNPQPLVNAQCTDDYYRGHSALHIAIEKRSLQCVK GHILILLGGIYLLVGQLWYFWRRHVFIWISFIDSYFEILFLFQALLTVVSQVLCFLAIEW gaseenyvpvollosn

(full-length predicted)

>hvR2.altFL

Fig. 16

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Agent: Mario Cloutier

Docket No.: MPI98-093P2RCP3DV1AM

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CLUSTAL W (1.74) multiple sequence alignment

CEOSIAE VV (1.7-	manufacture anguittern
humanVR2	MTSPSSSPVFRLETLDGGQEDGSEADRGKLDFGSGLPPMESQFQGEDRKFAPQIRVNLNY
hVR2.altFL	MTSPSSSPVFRLETLDGGQEDGSEADRGKLDFGSGLPPMESQFQGEDRKFAPQIRVNLNY
humanVR2	RKGTGASQPDPNRFDRDRLFNAVSRGVPEDLAGLPEYLSKTSKYLTDSEYTEGSTGKTCL
hVR2.altFL	RKGTGASQPDPNRFDRDRLFNAVSRGVPEDLAGLPEYLSKTSKYLTDSEYTEGSTGKTCL
humanVR2	MKAVLNLKDGVNACILPLLQIDRDSGNPQPLVNAQCTDDYYRGHSALHIAIEKRSLQCVK
hVR2.altFL	MKAVLNLKDGVNACILPLLQIDRDSGNPQPLVNAQCTDDYYRGHSALHIAIEKRSLQCVK
humanVR2	LLVENGANVHARACGRFFQKGQGTCFYFGELPLSLAACTKQWDVVSYLLENPHQPASLQA
hVR2.altFL	LLVENGANVHARACGRFFQKGQGTCFYFGELPLSLAACTKQWDVVSYLLENPHQPASLQA
humanVR2	TDSQGNTVLHALVMISDNSAENIALVTSMYDGLLQAGARLCPTVQLEDIRNLQDLTPLKL
hVR2.altFL	TDSQGNTVLHALVMISDNSAENIALVTSMYDGLLQAGARLCPTVQLEDIRNLQDLTPLKL
humanVR2	AAKEGKIEIFRHILQREFSGLSHLSRKFTEWCYGPVRVSLYDLASVDSCEENSVLEIIAF
hVR2.altFL	AAKEGKIEIFRHILQREFSGLSHLSRKFTEWCYGPVRVSLYDLASVDSCEENSVLEIIAF
humanVR2	HCKSPHRHRMVVLEPLNKLLQAKWDLLIPKFFLNFLCNLIYMFIFTAVAYHQPTLKKQAA
hVR2.altFL	HCKSPHRHRMVVLEPLNKLLQAKWDLLIPKFFLNFLCNLIYMFIFTAVAYHQPTLKKQAA
humanVR2	PHLKAEVGNSMLLTGHILILLGGIYLLVGQLWYFWRRHVFIWISFIDSYFEILFLFQALL
hVR2.altFL	PHLKAEVGNSMLLTGHILILLGGIYLLVGQLWYFWRRHVFIWISFIDSYFEILFLFQALL
humanVR2	TVVSQVLCFLAIEWYLPLLVSALVLGWLNLLYYTRGFQHTGIYSVMIQKVILRDLLRFLL
hVR2.altFL	TVVSQVLCFLAIEWYLPLLVSALVLGWLNLLYYTRGFQHTGIYSVMIQK
humanVR2 hVR2.altFL	IYLVFLFGFAVALVSLSQEAWRPEAPTGPNATESVQPMEGQEDEGNGAQYRGILEASLEL
humanVR2 hVR2.altFL	FKFTIGMGELAFQEQLHFRGMVLLLLLAYVLLTYILLLNMLIALMSETVNSVATDSWSIW
humanVR2 hVR2.altFL	KLQKAISVLEMENGYWWCRKKQRAGVMLTVGTKPDGSPDERWCFRVEEVNWASWEQTLPTKAISVLEMENGYWWCRKKQRAGVMLTVGTKPDGSPDERWCFRVEEVNWASWEQTLPT ***********************************
humanVR2 hVR2.altFL	LCEDPSGAGVPRTLENPVLASPPKEDEDGASEENYVPVQLLQSN LCEDPSGAGVPRTLENPVLASPPKEDEDGASEENYVPVQLLQSN ************************************

Fig. 17